

What is claimed is:

1. A method for ligating a first oligopeptide with a
5 second oligopeptide end to end for producing an
oligopeptide product, the method comprising the following
steps:

Step A: admixing the first and second oligopeptides in
a reaction solution including a catalytic thiol, the
10 first oligopeptide including a C-terminal thioester, the
second oligopeptide including an N-terminal cysteine
having an unoxidized sulfhydryl side chain; then

Step B: condensing the unoxidized sulfhydryl side
chain of the N-terminal cysteine with the C-terminal
15 thioester for producing an intermediate oligopeptide
linking the first and second oligopeptides with a β -
aminothioester bond; and then

Step C: rearranging the β -aminothioester bond of the
intermediate oligopeptide of said Step B for producing
20 the oligopeptide product linking the first and second
oligopeptides with an amide bond.

2. A method as described in Claim 1 wherein, in said
step A, the catalytic thiol is selected from the group
25 consisting of unconjugated mercaptans and conjugated
thiols.

3. A method as described in Claim 2 wherein, in said
step A, the catalytic thiol is benzyl mercaptan.

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4. A method as described in Claim 2 wherein, in said
step A, the catalytic thiol is a conjugated thiol
selected from the group consisting of thiophenol, 1-thio-
5 2-nitrophenol, 2-thio-benzoic acid, 2-thio-pyridine, 4-
thio-2-pyridinecarboxylic acid, and 4-thio-2-nitro-
pyridine.

5. A method as described in Claim 4 wherein, in said
10 step A, the conjugated thiol is thiophenol.

6. An oligopeptide intermediate comprising:
a first oligopeptide segment having a C-terminal
thioester,
15 a second oligopeptide segment having a N-terminal
cysteine, and
a β -aminothioester linkage unit linking the C-
terminal thioester and the N-terminal cysteine, said β -
aminothioester linkage unit spontaneously rearranging
20 intramolecularly to form an amide bond linking said first
and second oligopeptides segments end to end.

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7. A method for producing an oligopeptide having a C-terminal thioester, the method comprising the following steps:

5 Step A: providing a resin having a linker with an unoxidized thiol;

Step B: providing a Boc-amino acid succinimide ester; then

10 Step C:: admixing the resin of said Step A and the Boc-amino acid succinimide ester of said Step B under reaction conditions for producing a Boc-amino thioester-resin; then

15 Step D: assembling an oligopeptide onto the Boc-amino thioester-resin by stepwise solid phase peptide synthesis; then

Step E: cleaving the Boc-amino thioester-resin of said Step D with HS for producing an oligopeptide having a C-terminal thiol; and then

20 Step F: converting the oligopeptide having a C-terminal thiol of said Step E to the oligopeptide having a C-terminal thioester.

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